## **Neuroimaging Highlight**

Editors: Mark Hudon, Richard Farb

## Wallerian-Like Degeneration After Ischemic Stroke Revealed by Diffusion - Weighted Imaging

Submitted by: P.N. Sylaja, M. Goyal, T. Watson, M.D. Hill

Can. J. Neurol. Sci. 2007; 34: 243-244

A 22-year-old female was seen in the emergency within one hour of acute onset of right sided headache followed by weakness of the left side of body. On neurological examination, she was mildly drowsy, had forced right gaze deviation, dysarthria, left hemiplegia and left hemisensory loss. Computed tomography (CT) scan revealed early ischemic changes in the right middle cerebral artery (MCA) territory. The CT angiography done showed evidence of dissection of the supraclinoid segment of the right internal carotid artery with reduced flow distally into the MCA, which was confirmed by a conventional angiogram. In view of the intracranial carotid dissection, the patient was not treated with intravenous tissue plasminogen activator. Magnetic resonance imaging (MRI) of the brain done on the next day revealed evidence of acute ischemic lesions in the right MCA and anterior cerebral artery territory on diffusion-weighted imaging (DWI), with normal brainstem. [Figure 1] A repeat MRI performed 13 days after

ictus showed hyperintense signal on DWI in the right cerebral peduncle which was hypointense on apparent diffusion coefficient (ADC) map suggestive of Wallerian-like degeneration. [Figure 2] The signal changes were less conspicuous on T2-weighted images. She had antigravity strength in the left leg but remained weak in her left arm at one month.

Wallerian degeneration is defined as peripheral axonal degeneration after transection of a nerve trunk itself. In the central nervous system, Wallerian –like axonal degeneration occurs similarly; it is most frequently seen affecting the corticospinal tract after injury to the motor cortex. Wallerian-like degeneration can begin within one week of damage to the fiber tract and the demyelination can continue for the next six months. The availability of DWI and diffusion tensor-imaging (DTI) has facilitated early detection of Wallerian-like degeneration, within 2-14 days after ischemic stroke.<sup>2-4</sup>

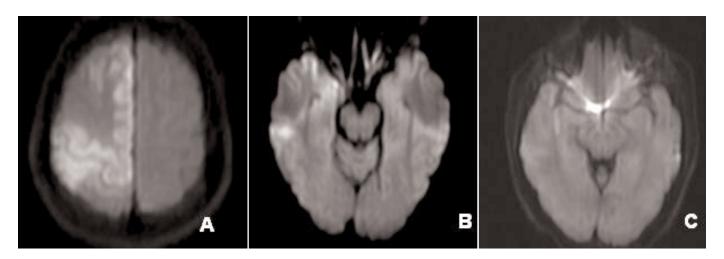


Figure 1: A. Diffusion-weighted imaging done on second day of stroke show acute ischemic lesions in the right frontoparietal regions corresponding to middle cerebral and anterior cerebral artery territory. B, C. No lesions are seen on the midbrain.

From the Calgary Stroke Program, Department of Clinical Neurosciences (PNS, TW, MDH), Department of Neuroradiology (MG), University of Calgary, Calgary, Alberta. Received September 18, 2006. Accepted in Final Form December 4, 2006.

Reprint requests to: Michael D. Hill, Calgary Stroke Program, Department of Clinical Neurosciences, Foothills Hospital, Room 1242A, Calgary, Alberta, T2N 2T9, Canada.

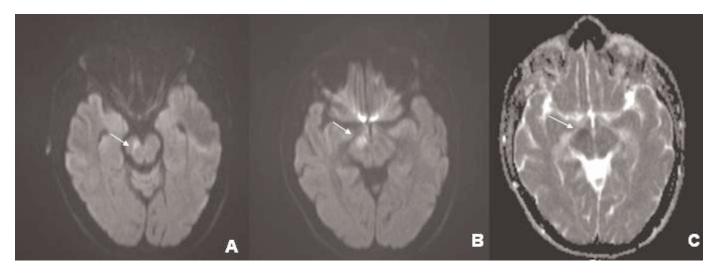


Figure 2: A & B. Diffusion weighted imaging shows increased signal intensity on the right cerebral peduncle corresponding to the corticospinal tract. C. The apparent diffusion coefficient map shows that the same area is hypointense.

Wallerian-like degeneration develops through four stages. The first stage is characterized by physical disintegration of the axons and myelin sheaths. This acute white matter tract injury is associated with cessation of energy dependant axo-plasmic transport and cytotoxic edema. The DWI identifies this earliest stage of Wallerian-like degeneration as a hyperintense signal which is hypointense on ADC map. But the conventional imaging will be normal during this stage. The second stage takes place 4-14 weeks after stroke and is characterized by rapid destruction of myelin and is seen as hypointense signal on T2 and proton density images. In the third stage gliosis occurs, which gives a hyperintense signal on T2 and fluid attenuated inversion recovery (FLAIR) sequences. The stage 4 or end stage occurs several years later and reveals volume loss from atrophy of the white matter tracts. Axonal degeneration of the middle cerebellar peduncle also has been reported following pontine infarction.<sup>5,6</sup> Diffusion tensor-imaging in particular the parallel and perpendicular diffusivity has been shown to reflect axon and myelin pathologies respectively in the optic nerve of a mouse model of retinal ischemia.7 Studies in humans have shown that, compared to DWI, DTI may be a better tool to stage Wallerianlike degeneration, as it correlates better with axonal and myelin integrity.<sup>3,8</sup> The exact prevalence of Wallerian-like degeneration after acute stroke is not known. A study by Castillo and Mukherji<sup>2</sup> detected evidence of Wallerian-like degeneration in 2 of the 11(18%) patients who under went DWI within 72 hours of ischemic stroke of the pyramidal tract after ischemic stroke reflects severe pyramidal tract damage and may slow the functional recovery and has been associated with persisting impairment of motor functions, as was observed in our patient.9-11 Further prospective studies are needed to establish the correlation of the DWI changes of Wallerian-like degeneration and the functional outcome to determine its utility as a prognostic marker after stroke. Identification of these lesions is important, as these new lesions on follow-up imaging should not be misinterpreted as new ischemic lesions.

## REFERENCES

- Kuhn MJ, Mikulis DJ, Ayoub DM, Kosofsky BE, Davis KR, Taveras JM.Wallerian degeneration after cerebral infarction: evaluation with sequential MR imaging. Radiology. 1989;172: 179-82.
- Castillo M, Mukherji SK. Early abnormalities related to postinfarction Wallerian degeneration with MR diffusion-weighted imaging. J Comput Assist Tomogr. 1999;23:1004-7.
- Thomalla G, Glauche V, Koch MA, Beaulieu C, Weiller C, Rother J. Diffusion tensor imaging detects early Wallerian degeneration of the pyramidal tract after ischemic stroke. Neuroimage. 2004;22:1767-74.
- Uchino A, Sawada A, Takase Y, Egashira R, Kudo S. Transient detection of early Wallerian degeneration on diffusion-weighted MRI after an acute cerebrovascular accident. Neuroradiology. 2004;46:183-8.
- Kuker W, Schmidt F, Heckl S, Nagele T, Herrlinger U. Bilateral Wallerian degeneration of the middle cerebellar peduncles due to paramedian pontine infarction:MRI findings. Neuroradiology. 2004;46:896-9.
- Fujii A, Tokuda A, Yoneda M, Kuriyama M. A case showing Wallerian degeneration of the bilateral middle cerebellar peduncles on MRI followed by the right pontine infarction Rinsho Shinkeigaku. 2004;44:105-7.
- Song SK, Sun SW, Ju WK, Lin SJ, Cross AH, Neufeld AH.
  Diffusion tensor imaging detects and differentiates axon and
  myelin degeneration in mouse optic nerve after retinal ischemia.
  Neuroimage. 2003;20:1714-22.
- Concha L, Gross DW, Wheatley BM, Beaulieu C. Diffusion tensor imaging of time dependant axonal and myelin degradation aftercorpus callosotomy in epilepsy patients. Neuroimage. 2006;32:1090-9.
- Watanabe T, Honda Y, Fujii Y, Koyama M, Matsuzawa H, Tanaka R. Three dimensional anisotropy contrast magnetic resonance axonography to predict the prognosis for motor function in patients suffering from stroke. J Neurosurg. 2001;94:955-60.
- Miyai I, Suzuki T, Kii K, Kang J, Kubota K. Wallerian degeneration of the pyramidal tract does not affect stroke rehabilitation outcome. Neurology. 1998;51:1613-6.
- Mazumdar A, Mukherjee P, Miller JH, Malde H, McKinstry RC. Diffusion-weighted imaging of acute corticospinal tract injury preceding Wallerian degeneration in the maturing human brain. AJNR. 2003;24:1057-66.